I

CLAIMS

1. An amide derivative of the Formula I

a'

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$$(R^2)_p$$
 H
 $(CH_2)_q$
 R^4

wherein R3 is (1-6C)alkyl or halogeno;

Q is aryl or heteroaryl which optionally bears 1, 2, 3 or 4 substituents selected from hydroxy, halogeno, trifluoromethyl, cyano, mercapto, nitro, amino, carboxy, carbamoyl, formyl, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (1-3C)alkylenedioxy,

- 10 (1-6C)alkylthio, (1-6C)alkylsulphinyl, (1-6C)alkylsulphonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N-(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (1-6C)alkanoylamino, N-(1-6C)alkylsulphamoyl, N-(1-6C)alkyl]sulphamoyl, (1-6C)alkanoylamino, N-(1-6C)alkyl-(1-6C)alkyl-(1-6C)alkyl, hydroxy-(1-6C)alkyl,
- 15 (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl, di-[(1-6C)alkyl]amino-(1-6C)alkyl, carboxy-(1-6C)alkyl, (1-6C)alkoxycarbonyl-(1-6C)alkyl, carbamoyl-(1-6C)alkyl, N-(1-6C)alkylcarbamoyl-(1-6C)alkyl, NN-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkyl, halogeno-(2-6C)alkoxy, hydroxy-(2-6C)alkoxy, (1-6C)alkoxy, cyano-(1-6C)alkoxy, carboxy-(1-6C)alkoxy,
- 20 (1-6C)alkoxycarbonyl-(1-6C)alkoxy, carbamoyl-(1-6C)alkoxy, N-(1-6C)alkylcarbamoyl-(1-6C)alkoxy, N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkoxy, amino-(2-6C)alkoxy, (1-6C)alkylamino-(2-6C)alkoxy, di-[(1-6C)alkyl]amino-(2-6C)alkoxy, halogeno-(2-6C)alkylamino, hydroxy-(2-6C)alkylamino, (1-6C)alkoxy-(2-6C)alkylamino, cyano-(1-6C)alkylamino, carboxy-(1-6C)alkylamino, (1-6C)alkoxycarbonyl-(1-6C)alkylamino,
- 25 carbamoyl-(1-6C)alkylamino, N-(1-6C)alkylamino, N-(1-6C)alkylamino, N-(1-6C)alkylamino, amino-(2-6C)alkylamino, (1-6C)alkylamino-(2-6C)alkylamino, di-[(1-6C)alkylamino-(2-6C)alkylamino, di-[(1-6C)alkyl]amino-(2-6C)alkylamino,



 \underline{N} -(1-6C)alkyl-halogeno-(1-6C)alkylamino, \underline{N} -(1-6C)alkyl-hydroxy-(2-6C)alkylamino, \underline{N} -(1-6C)alkyl-(1-6C)alkoxy-(2-6C)alkylamino, \underline{N} -(1-6C)alkyl-cyano-(1-6C)alkylamino, \underline{N} -(1-6C)alkyl-carboxy-(1-6C)alkylamino, \underline{N} -(1-6C)alkyl-(1-6C)alkylamino, \underline{N} -(1-6C)alkylamino, \underline{N} -(1-6C)alkyl-carbamoyl-(1-6C)alkylamino, \underline{N} -(1-6C)alkyl-

- 5 N-(1-6C)alkylcarbamoyl-(1-6C)alkylamino, N-(1-6C)alkyl-N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkylamino, N-(1-6C)alkyl-amino-(2-6C)alkylamino, N-(1-6C)alkyl-di-[(1-6C)alkyl]amino-(2-6C)alkylamino, N-(1-6C)alkyl-di-[(1-6C)alkyl]amino-(2-6C)alkylamino, halogeno-(2-6C)alkanoylamino, hydroxy-(2-6C)alkanoylamino, (1-6C)alkoxy-(2-6C)alkanoylamino, cyano-(2-6C)alkanoylamino, carboxy-(2-6C)alkanoylamino,
- 10 (1-6C)alkoxycarbonyl-(2-6C)alkanoylamino, carbamoyl-(2-6C)alkanoylamino, N-(1-6C)alkylcarbamoyl-(2-6C)alkanoylamino, N-(1-6C)alkylcarbamoyl-(2-6C)alkanoylamino, (1-6C)alkylamino-(2-6C)alkanoylamino, di-[(1-6C)alkyl]amino-(2-6C)alkanoylamino, aryl, aryl-(1-6C)alkyl, aryl-(1-6C)alkoxy, aryloxy, arylamino, N-(1-6C)alkyl-arylamino, aryl-(1-6C)alkylamino, N-(1-6C)alkyl-
- 15 aryl-(1-6C)alkylamino, aroylamino, arylsulphonylamino, N-arylsulphamoyl, aryl-(2-6C)alkanoylamino, heteroaryl, heteroaryl-(1-6C)alkyl, heteroaryloxy, heteroaryl-(1-6C)alkoxy, heteroarylamino, N-(1-6C)alkyl-heteroarylamino, heteroaryl-(1-6C)alkylamino, heteroaryl-(1-6C)alkylamino, heteroarylsulphonylamino, N-heteroarylsulphamoyl, heteroaryl-(2-6C)alkanoylamino,
- heterocyclyl, heterocyclyl-(1-6C)alkyl, heterocyclyloxy, heterocyclyl-(1-6C)alkoxy, heterocyclylamino, N-(1-6C)alkyl-heterocyclylamino, heterocyclyl-(1-6C)alkylamino, N-(1-6C)alkyl-heterocyclyl-(1-6C)alkylamino, heterocyclylcarbonylamino, heterocyclylsulphonylamino, N-heterocyclylsulphamoyl and heterocyclyl-(2-6C)alkanoylamino,
- and wherein any of the substituents on Q defined hereinbefore which comprise a CH₂ group which is attached to 2 carbon atoms or a CH₃ group which is attached to a carbon atom may optionally bear on each said CH₂ or CH₃ group a substituent selected from hydroxy, amino, (1-6C)alkoxy, (1-6C)alkylamino, di-[(1-6C)alkyl]amino and heterocyclyl; and wherein any aryl, heteroaryl or heterocyclyl group in a substituent on Q may optionally
- bear 1 or 2 substituents selected from hydroxy, halogeno, (1-6C)alkyl, (1-6C)alkoxy, carboxy, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N-(1-6C)alkylcarbamoyl, N-(1-6C)alkylcarbamoyl,

(2-6C)alkanoyl, amino, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkyl, di-[(1-6C)alkyl]amino-(1-6C)alkyl, aryl and aryl-(1-6C)alkyl; R² is hydroxy, halogeno, trifluoromethyl, cyano, mercapto, nitro, amino, carboxy,

5 (1-6C)alkoxycarbonyl, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (1-6C)alkylamino or di-[(1-6C)alkyl]amino;

p is 0, 1 or 2;

q is 0, 1, 2, 3 or 4; and

R⁴ is aryl, aryl-(1-6C)alkoxy, aryloxy, arylamino, N-(1-6C)alkyl-arylamino,

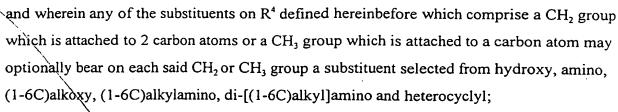
- aryl-(1-6C)alkylamino, N-arylsulphamoyl, aryl-(2-6C)alkylamino, aroylamino, cycloalkyl, heteroaryl, heteroaryloxy, heteroaryl-(1-6C)alkoxy, heteroarylamino, N-(1-6C)alkyl-heteroarylamino, heteroaryl-(1-6C)alkylamino, N-(1-6C)alkyl-heteroaryl-(1-6C)alkylamino, heteroarylsulphonylamino, N-heteroarylsulphamoyl, heteroaryl-
- 15 (2-6C)alkanoylamino, heterocyclyl, heterocyclyloxy, heterocyclyl-(1-6C)alkoxy, heterocyclylamino, N-(1-6C)alkyl-heterocyclylamino, heterocyclyl-(1-6C)alkylamino, N-(1-6C)alkyl-heterocyclyl-(1-6C)alkylamino, heterocyclylcarbonylamino, heterocyclylsulphonylamino, N-heterocyclylsulphonylamino or heterocyclyl-(2-6C)alkanoylamino and R⁴ optionally bears 1, 2, 3 or 4 substituents selected from hydroxy, halogeno,
- 20 trifluoromethyl, cyano, mercapto, nitro, amino, carbòxy, carbamoyl, formyl, (1-6C)alkyl, (2-6C)alkenyl, (2-6C)alkynyl, (1-6C)alkoxy, (1-3C)alkylenedioxy, (1-6C)alkylthio, (1-6C)alkylsulphinyl, (1-6C)alkylsulphonyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, NN-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, (2-6C)alkanoyloxy, (1-6C)alkanoylamino, N-(1-6C)alkylsulphamoyl,
- 25 N,N-di-[(1-6C)alkyl]sulphamoyl, (1-6C)alkanesulphonylamino, N-(1-6C)alkyl-(1-6C)alkyl-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl, di-[(1-6C)alkyl]amino-(1-6C)alkyl, carboxy-(1-6C)alkyl, (1-6C)alkyl, (1-6C)alkyl, carbamoyl-(1-6C)alkyl, N-(1-6C)alkyl, N-(1-6C)alkyl, not carbamoyl-(1-6C)alkyl, N-(1-6C)alkyl, N-(1-6C)alkyl, not carbamoyl-(1-6C)alkyl, N-(1-6C)alkyl, not carbamoyl-(1-6C)alkyl, no
- 30 N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkyl, halogeno-(2-6C)alkoxy, hydroxy-(2-6C)alkoxy, (1-6C)alkoxy, cyano-(1-6C)alkoxy, carboxy-(1-6C)alkoxy,

a' Cont

-6C)alkoxycarbonyl-(1-6C)alkoxy, carbamoyl-(1-6C)alkoxy, N-(1-6C)alkylcarbamoyl-(1-8C)alkoxy, N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkoxy, amino-(2-6C)alkoxy, (1-6C)alkylamino-(2-6C)alkoxy, di-[(1-6C)alkyl]amino-(2-6C)alkoxy, halogeno-(2-6C)alkylamino, hydroxy-(2-6C)alkylamino, (1-6C)alkoxy-(2-6C)alkylamino, cvano-5 (1-6C)alkylamino, carboxy-(1-6C)alkylamino, (1-6C)alkoxycarbonyl-(1-6C)alkylamino. carbamoyl-(1-6C)alkylamino, N-(1-6C)alkylcarbamoyl-(1-6C)alkylamino, N,N-di-[(1-6C)alky\carbamoyl-(1-6C)alkylamino, amino-(2-6C)alkylamino, (1-6C)alkylamino-(2-6C)alkylamino, di-[(1-6C)alkyl]amino-(2-6C)alkylamino, N-(1-6C)alkyl-halogeno-(1-6C)alkylamino, N-(1-6C)alkyl-hydroxy-(2-6C)alkylamino, 10 N-(1-6C)alkyl-(1-6C)alkoxy-(2-6C)alkylamino, N-(1-6C)alkyl-cyano-(1-6C)alkylamino, N-(1-6C)alkyl-carboxy-(1-6C)alkylamino, N-(1-6C)alkyl-(1-6C)alkoxycarbonyl-(1-6C)alkylamino, N-(1-6C)alkyl-carbamoyl-(1-6C)alkylamino, N-(1-6C)alkyl-N-(1-6C)alkylcarbamoyl-(1-6C)alkylamino, N-(1-6C)alkyl-N,N-di-[(1-6C)alkyl]carbamoyl-(1-6C)alkylamino, N-(1-6C)alkyl-amino-(2-6C)alkylamino, N-(1-6C)alkyl-(1-6C)alkylamino-15 (2-6C)alkylamino, N-(1-6C)alkyl-di-[(1-6C)alkyl]amino-(2-6C)alkylamino, halogeno-(2-6C)alkanoylamino, hydroxy-(2-6C)alkanoylamino, (1-6C)alkoxy-(2-6C)alkanoylamino, cyano-(2-6C)alkanoylamino, carboxy-(2-6C)alkanoylamino, (1-6C)alkoxycarbonyl-(2-6C)alkanoylamino, carbamoyl-(2-6C)alkanoylamino, N-(1-6C)alkylcarbamoyl-(2-6C)alkanoylamino, N,N-di-[(1-6C)alkyl]carbamoyl-(2-6C)alkanoylamino,

- 20 amino-(2-6C)alkanoylamino, (1-6C)alkylamino-(2-6C)alkanoylamino, di-[(1-6C)alkyl]amino-(2-6C)alkanoylamino, aryl, aryl-(1-6C)alkyl, aryl-(1-6C)alkoxy, aryloxy, arylamino, N-(1-6C)alkyl-arylamino, aryl-(1-6C)alkylamino, N-(1-6C)alkyl-aryl-(1-6C)alkylamino, arylsulphonylamino, N-arylsulphamoyl, aryl-(2-6C)alkanoylamino, heteroaryl, heteroaryl-(1-6C)alkyl, heteroaryloxy, heteroaryl-(1-6C)alkoxy, heteroarylamino,
- 25 <u>N</u>-(1-6C)alkyl-heteroarylamino, heteroaryl-(1-6C)alkylamino, <u>N</u>-(1-6C)alkyl-heteroaryl-(1-6C)alkylamino, heteroarylsulphonylamino, <u>N</u>-heteroarylsulphamoyl, heteroaryl-(2-6C)alkanoylamino, heterocyclyl, heterocyclyl-(1-6C)alkyl, heterocyclyloxy, heterocyclyl-(1-6C)alkoxy, heterocyclylamino, <u>N</u>-(1-6C)alkyl-heterocyclyl-heterocyclyl-(1-6C)alkyl-heterocyclyl-
- 30 (1-6C)alkylamino, heterocyclylcarbonylamino, heterocyclylsulphonylamino, N-heterocyclylsulphamoyl and heterocyclyl-(2-6C)alkanoylamino,





- and wherein any aryl, heteroaryl or heterocyclyl group in a substituent on R⁴ may optionally bear 1 or 2 substituents selected from hydroxy, halogeno, (1-6C)alkyl, (1-6C)alkoxy, carboxy, (1-6C)alkoxycarbonyl, N-(1-6C)alkylcarbamoyl, N,N-di-[(1-6C)alkyl]carbamoyl, (2-6C)alkanoyl, amino, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, halogeno-(1-6C)alkyl, hydroxy-(1-6C)alkyl, (1-6C)alkoxy-(1-6C)alkyl, cyano-(1-6C)alkyl, amino-(1-6C)alkyl,
- 10 (1-6C)alkylamino-(1-6C)alkyl di-[(1-6C)alkyl]amino-(1-6C)alkyl, aryl and aryl-(1-6C)alkyl; or a pharmaceutically-acceptable salt or <u>in-vivo</u>-cleavable ester thereof; except that the compounds:-
 - N-(2-cyclohexylethyl)-3-(4-hydroxybenzamido)-4-methylbenzamide,
 - 3-(4-aminobenzamido)-N-(4-carboxy-3-hydroxyphenyl)-4-methylbenzamide,
- 15 N-(4-carboxy-3-hydroxyphenyl)-4-methyl-3-(4-nitrobenzamido)benzamide,
 - 3-(4-aminobenzamido)-4-methyl-N-(2-pyridyl)benzamide,
 - 4-methyl-3-(4-nitrobenzamido)-N-(2-pyridyl)benzamide,
 - $3-(4-aminobenzamido)-4-methyl-\underline{N}-(2-thiazolyl) benzamide,$
 - 4-methyl-3-(4-nitrobenzamido)-N-(2-thiazolyl)benzamide,
- 20 3-benzamido-4-chloro- \underline{N} -(2-fluoroanilino)benzamide,
 - 3-(2-hydroxy-4-methylbenzamido)-N-(4-hydroxyphenyl)-4-methylbenzamide,
 - 3-(3-hydroxy-2-naphthoylamino)-4-methyl-N-phenylbenzamide and
 - 4-chloro-3-(3-hydroxy-2-naphthoylamino)-2-methyl-N-phenylbenzamide are excluded.
- 25 2. An amide derivative of the Formula I according to claim 1 wherein R³ is methyl, ethyl, chloro or bromo;

Q is phenyl which bears 1, 2 or 3 substituents selected from hydroxy, fluoro, chloro, trifluoromethyl, cyano, carboxy, methyl, ethyl, propyl, methoxy, ethoxy, methylenedioxy, methoxycarbonyl, ethoxycarbonyl, tert-butoxycarbonyl, acetyl, propionyl, chloromethyl,

30 methoxymethyl, methylaminomethyl, ethylaminomethyl, dimethylaminomethyl, diethylaminomethyl, 2-chloroethoxy, 3-chloropxopoxy, 2-hydroxyethoxy, 3-hydroxypropoxy,



2-methoxyethoxy, 2-ethoxyethoxy, 3-methoxypropoxy, 3-ethoxypropoxy, cyanomethoxy, carboxymethoxy, methoxycarbonylmethoxy, ethoxycarbonylmethoxy,

tert-butdxycarbonylmethoxy, 2-aminoethoxy, 3-aminopropoxy, 2-methylaminoethoxy,

- 2-ethylaminoethoxy, 3-methylaminopropoxy, 3-ethylaminopropoxy,
- 5 2-dimethylaminoethoxy, 2-diethylaminoethoxy, 3-dimethylaminopropoxy,
 - 3-diethylaminopropoxy, 2-pyridylmethoxy, 2-(imidazol-1-yl)ethoxy,
 - 3-(imidazol-1-yl)propoxy, pyrrolidin-1-yl, piperidino, morpholino, piperazin-1-yl,
 - 4-methylpiperazin-lyl, 4-acetylpiperazin-l-yl, pyrrolidin-l-ylmethyl, piperidinomethyl, morpholinomethyl, piperazin-l-ylmethyl, 4-methylpiperazin-l-ylmethyl, 4-acetylpiperazin-
- 10 1-ylmethyl, piperidin-4-yloxy, 1-methylpiperidin-4-yloxy, 2-(pyrrolidin-1-yl)ethoxy, 3-(pyrrolidin-1-yl)propoxy, 2-piperidinoethoxy, 3-piperidinopropoxy, 2-morpholinoethoxy, 3-morpholinopropoxy, 2-piperazin-1-ylethoxy, 3-piperazin-1-ylpropoxy, 2-(4-methylpiperazin-1-yl)ethoxy, 3-(4-methylpiperazin-1-yl)propoxy, 2-(4-acetylpiperazin-1-yl)ethoxy and 3-(4-acetylpiperazin-1-yl)propoxy,
- or Q is furyl, thienyl, oxazolyl, isoxazolyl, imidazolyl, thiazolyl, isothiazolyl, pyridyl, pyridazinyl, pyrimidinyl, pyrazinyl, benzofuranyl, indolyl, benzothienyl, benzoxazolyl, benzimidazolyl, benzothiazolyl, indazolyl, quinolyl, isoquinolyl, quinazolinyl, quinoxalinyl or naphthyridinyl which optionally bears 1 or 2 substituents selected from hydroxy, fluoro, chloro, trifluoromethyl, cyano, methyl, ethyl, methoxy and ethoxy;
- 20 p is 0;

q is 0; and

R⁴ is phenyl which bears 1 or 2 substituents selected from hydroxy, fluoro, chloro, trifluoromethyl, cyano, amino, methyl, ethyl, methoxy, ethoxy, methylenedioxy, methylamino, ethylamino, dimethylamino, diethylamino, acetyl, propionyl, chloromethyl,

- 25 methoxymethyl, 2-methoxyethyl, methylaminomethyl, ethylaminomethyl, dimethylaminomethyl, diethylaminomethyl, 2-chloroethoxy, 3-chloropropoxy, 2-hydroxyethoxy, 3-hydroxypropoxy, 2-methoxyethoxy, 2-ethoxyethoxy, 3-methoxypropoxy, 3-ethoxypropoxy, cyanomethoxy, carboxymethoxy, methoxycarbonylmethoxy, ethoxycarbonylmethoxy, tert-butoxycarbonylmethoxy, 2-aminoethoxy, 3-aminopropoxy,
- 30 2-methylaminoethoxy, 2-ethylaminoethoxy, 3-methylaminopropoxy, 3-ethylaminopropoxy, 2-dimethylaminoethoxy, 2-diethylaminoethoxy, 3-dimethylaminopropoxy,

ī D 3-diethylaminopropoxy, 2-chloroethylamino, 2-hydroxyethylamino,

2-methoxyethylamino, 2-ethoxyethylamino, 2-aminoethylamino,

2-meth) laminoethylamino, 2-ethylaminoethylamino, 2-dimethylaminoethylamino,

2-diethylamino, \underline{N} -(2-chloroethyl)- \underline{N} -methylamino, \underline{N} -(2-hydroxyethyl)-

5 <u>N</u>-methylamino, <u>N</u>-(2-methoxyethyl)-<u>N</u>-methylamino, <u>N</u>-(2-ethoxyethyl)-

 \underline{N} -methylamino, \underline{N} -(2-aminoethyl)- \underline{N} -methylamino, \underline{N} -(2-methylaminoethyl)-

 \underline{N} -methylamino, \underline{N} -(2-dimethylaminoethyl)- \underline{N} -methylamino, \underline{N} -(3-aminopropyl)-

 \underline{N} -methylamino, \underline{N} -(3-ethylaminopropyl)- \underline{N} -methylamino, \underline{N} -(3-ethylaminopropyl)-

 \underline{N} -methylamino, \underline{N} -(3-dimethylaminopropyl)- \underline{N} -methylamino, \underline{N} -(3-diethylaminopropyl)-

- N-methylamino, phenyl, benzyl, benzyloxy, 2-pyridylmethoxy, 2-(imidazol-1-yl)ethoxy, 3-(imidazol-1-yl)propoxy, pyrrolidin-1-yl, piperidino, morpholino, piperazin-1-yl, 4-methylpiperazin-1-yl, 4-acetylpiperazin-1-yl, pyrrolidin-1-ylmethyl, piperidinomethyl, morpholinomethyl, piperazin-1-ylmethyl, 4-methylpiperazin-1-ylmethyl, 4-acetylpiperazin-1-ylmethyl, piperidin-4-yloxy, 1-methylpiperidin-4-yloxy, 2-(pyrrolidin-1-yl)ethoxy,
- 15 3-(pyrrolidin-1-yl)propoxy, 2-piperidinoethoxy, 3-piperidinopropoxy,
 - 2-morpholinoethoxy, 3-morpholinopropoxy, 2-piperazin-1-ylethoxy,
 - 3-piperazin-1-ylpropoxy, 2-(4-methylpiperazin-1-yl)ethoxy,
 - 3-(4-methylpiperazin-1-yl)propoxy, 2-(4-acetylpiperazin-1-yl)ethoxy and
 - 3-(4-acetylpiperazin-1-yl)propoxy;
- or a pharmaceutically-acceptable salt thereof; except that 3-(2-hydroxy-4-methylbenzamido)-N-(4-hydroxyphenyl)-4-methylbenzamide is excluded.
 - 3. An amide derivative of the Formula I according to claim 1
- 25 wherein R3 is methyl or chloro;

Q is phenyl which bears 1, 2 or 3 substituents selected from hydroxy, cyano, carboxy, methyl, ethyl, propyl, methoxy, acetyl and 2-methoxyethoxy;

p is 0;

q is 0; and

30 R⁴ is phenyl which bears 1 or 2 substituents selected from chloro, cyano and dimethylamino; or a pharmaceutically-acceptable salt thereof.

4. An amide derivative of the Formula I according to claim 1 wherein R³ is methyl or chloro;

Q is 3-isoxazolyl, 3-pyridyl or 6-quinolyl which optionally bears a substituent selected from chloro and methyl:

5 p is 0;

q is 0; and

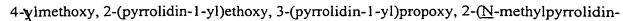
R⁴ is phenyl which bears a dimethylamino substituent; or a pharmaceutically-acceptable salt thereof.

- An amide derivative of the Formula I according to claim 1 wherein Q is substituted by a basic substituent selected from the substituents for Q defined in claim 1 and R⁴ is a phenyl or heteroaryl group as defined in claim 1 which also bears a basic substituent selected from the substituents for R⁴ defined in claim 1.
- 15 6. An amide derivative of the Formula I according to claim 1 wherein R³ is methyl or chloro;

Q is phenyl which bears a substituent selected from dimethylaminomethyl, diethylaminomethyl, \underline{N} -butyl- \underline{N} -methylaminomethyl, 2-dimethylaminoethoxy,

- 2-diethylaminoethoxy, 2-diisopropylaminoethoxy, 3-dimethylaminopropoxy,
- 3-diethylaminopropoxy, 3-diisopropylaminopropoxy, pyrrolidin-1-ylmethyl, 3-hydroxypyrrolidin-1-ylmethyl, morpholinomethyl, piperidinomethyl, homopiperidinomethyl, piperazin-1-ylmethyl, homopiperazin-1-ylmethyl, 4-methylpiperazin-1-ylmethyl, 4-ethylpiperazin-1-ylmethyl, 4-ethylpiperazin-1-ylmethyl, 4-ethylpiperazin-1-ylmethyl,
- 25 4-(2-hydroxyethyl)piperazin-1-ylmethyl, 2-pyridylmethoxy, pyrrolidin-3-yloxy, 1-methylpyrrolidin-3-yloxy, piperidin-3-yloxy, 1-methylpiperidin-3-yloxy, homopiperidin-3-yloxy, piperidin-4-yloxy, 1-methylpiperidin-4-yloxy, homopiperidin-4-yloxy, pyrrolidin-3-ylmethoxy, 1-methylpyrrolidin-3-ylmethoxy, piperidin-3-ylmethoxy, 1-methylpiperidin-3-ylmethoxy,
- 30 homopiperidin-3-ylmethoxy, 1-methylhomopiperidin-3-ylmethoxy, piperidin-4-ylmethoxy, 1-methylpiperidin-4-ylmethoxy, homopiperidin-4-ylmethoxy, 1-methylhomopiperidin-





- 2-yl)ethoxy, 3-(N-methylpyrrolidin-2-yl)propoxy, 2-piperidinoethoxy, 3-piperidinopropoxy,
- 2-morpholinoethoxy, 3-morpholinopropoxy, 2-piperazin-1-ylethoxy, 2-homopiperazin-
- 1-ylethoxy, 3-piperazin-1-ylpropoxy, 3-homopiperazin-1-ylpropoxy, 2-(4-methylpiperazin-
- 5 1-yl)ethoxy, 2-(4-methylhomopiperazin-1-yl)ethoxy, 3-(4-methylpiperazin-1-yl)propoxy,
 - 3-(4-methylhomopiperazin-1-yl)propoxy, 2-(4-acetylpiperazin-1-yl)ethoxy,
 - 3-(4-acetylpiperazin-1-yl)propoxy, 2-methoxyethylaminomethyl,
 - 3-methoxypropylaminomethyl, 2-aminoethylaminomethyl, 3-aminopropylaminomethyl,
 - 3-dimethylamino-2,2-dimethylpropylaminomethyl, 2-methylaminoethylaminomethyl,
- 10 3-methylaminopropylaminomethyl, 2-dimethylaminoethylaminomethyl,
 - 3-dimethylaminopropylaminomethyl, N-(2-methylaminoethyl)-N-methylaminomethyl,
 - \underline{N} -(3-methylaminopropyl)- \underline{N} -methylaminomethyl, \underline{N} -(2-dimethylaminoethyl)-
 - N-methylaminomethyl, N-(3-dimethylaminopropyl)-N-methylaminomethyl and
 - 3-morpholinopropylaminomethy, and Q is optionally substituted with a further substituent
- 15 selected from methyl and methoxy

p is 0;

q is 0; and

R⁴ is phenyl which is substituted at the 3 position with a substituent selected from dimethylamino, diethylamino, pyrrolidin-1 yl, piperidino, morpholino, piperazin-1-yl,

- 20 homopiperazin-1-yl, 4-methylpiperazin-1-yl and 4-methylhomopiperazin-1-yl and R⁴ is optionally substituted with a further substituent selected from fluoro, chloro, cyano, methyl and trifluoromethyl;
 - or a pharmaceutically-acceptable salt thereof.
- 25 7. An amide derivative of the Formula I according to claim 1 wherein R³ is methyl or chloro;
 - Q is 3-pyridyl or 4-pyridyl which bears a substituent selected from 2-aminoethylamino,
 - 3-aminopropylamino, 2-amino-2-methylpropylamino, 4-aminobutylamino,
 - 2-methylaminoethylamino, 2-ethylamino, 3-methylaminopropylamino,
- 30 4-methylaminobutylamino, 2-dimethylaminoethylamino, 2-diethylamino,
 - 3-dimethylaminopropylamino, 4-dimethylaminobutylamino, N-(2-methylaminoethyl)-

N-methylamino, N-(3-methylaminopropyl)-N-methylamino, N-(4-methylaminobutyl)-N-methylamino, N-(2-dimethylaminoethyl)-N-methylamino, N-(3-dimethylaminopropyl)-N-methylamino, N-(4-dimethylaminobutyl)-N-methylamino, pyrrolidin-1-yl, 3-hydroxypyrrolidin-1-yl, morpholino, piperidino, homopiperidino, piperazin-1-yl, 4-methylpiperazin-1-yl, 4-ethylpiperazin-1-yl, 4-(2-hydroxyethyl)piperazin-1-yl, 4-methylhomopiperazin-1-yl, 3-morpholinopropylamino or 2-(1-methylpyrrolidin-2-yl)ethylamino; p is 0;

q is 0; and

- 10 R⁴ is phenyl which is substituted at the 3-position with a substituent selected from dimethylamino, diethylamino, pyrrolidin-1-yl, piperidino, morpholino, piperazin-1-yl, homopiperazin-1-yl, 4-methylpiperazin-1-yl and 4-methylhomopiperazin-1-yl and R⁴ is optionally substituted with a further substituent selected from fluoro, chloro, cyano, methyl and trifluoromethyl;
- 15 or a pharmaceutically-acceptable salt thereof.
 - 8. An amide derivative of the Formula I according to claim 1 selected from:

 N-(3-dimethylaminophenyl)-4-methyl-3-(4-propylbenzamido)benzamide,

 3-(3,4-dimethoxybenzamido)-N-(3-dimethylaminophenyl)-4-methylbenzamide,
- 3-(4-butoxybenzamido)-N-(3-dimethylaminophenyl)-4-methylbenzamide, 4-chloro-N-(3-dimethylaminophenyl)-3-(4-propylbenzamido)benzamide, 3-(4-carboxybenzamido)-N-(3-dimethylaminophenyl)-4-methylbenzamide, N-(3,4-dichlorobenzyl)-3-(3,4,5-trimethoxybenzamido)-4-methylbenzamide, N-(2-cyclohexylethyl)-3-(3,4-dimethoxybenzamido)-4-methylbenzamide,
- 25 N-(3-dimethylaminophenyl)-4-methyl-3-(6-quinolylcarbonylamino)benzamide, 4-chloro-N-(3-dimethylaminophenyl)-3-(6-quinolylcarbonylamino)benzamide, 4-methyl-N-(3-morpholinophenyl)-3-(3-piperidin-4-yloxybenzamido)benzamide, 4-chloro-N-(3-fluoro-5-morpholinophenyl)-3-[3-(1-methylhomopiperidin-4-yloxy)benzamido]benzamide,
- 30 3-(2-diisopropylaminoethoxybenzamido)-4-methyl-<u>N</u>-(3-morpholinophenyl)benzamide, 3-(4-diethylaminomethylbenzamido)-4-methyl-<u>N</u>-(3-morpholinophenyl)benzamide,

Cont

4-methyl-3-[3-(4-methylhomopiperazin-1-ylmethyl)benzamido]-N-(3-morpholinophenyl)-benzamide

4-methyl-3-[3-(4-methylpiperazin-1-ylmethyl)benzamido]-N-(3-morpholinophenyl)-benzamide and

3-[6-(2-amino-2-methylpropylamino)pyrid-3-ylcarbonylamino]-4-chloro-N-(3-fluoro-5-morpholinophenyl)benzamide; or a pharmaceutically-acceptable salt thereof.

- A process for the preparation of an amide derivative of the Formula I, or a
 pharmaceutically-acceptable salt or <u>in-vivo</u>-cleavable ester thereof, according to claim 1 which comprises:
 - (a) reacting a benzoic acid of the Formula II, or a reactive derivative thereof,

with an amine of the Formula III

$$H_2N \longrightarrow (CH_2)_q \longrightarrow R^4$$

III

under standard amide bond forming conditions, wherein variable groups are as defined in claim 1 and wherein any functional group is protected if necessary, and:

- (i) removing any protecting groups; and
- (ii) optionally forming a pharmaceutically-acceptable salt or <u>in-vivo</u>-cleavable ester;
- (b) reacting an acid of the Formula IV, or an activated derivative thereof,

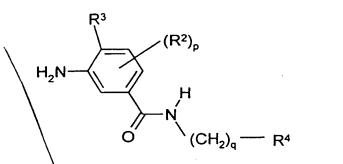
with an aniline of the Formula VI

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under standard amide bond forming conditions as defined hereinbefore, wherein variable groups are as defined in claim 1 and wherein any functional group is protected, if necessary, and:

- (i) removing any protecting groups;
 - (ii) optionally forming a pharmaceutically-acceptable salt or <u>in-vivo</u>-cleavable ester;
- (c) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is (1-6C)alkoxy or substituted (1-6C)alkoxy, (1-6C)alkylthio, (1-6C)alkylamino,
- 10 di-[(1-6C)alkyl]amino or substituted (1-6C)alkylamino or heterocyclyloxy, the alkylation, conveniently in the presence of a suitable base, of an amide derivative of the Formula I wherein a substituent on Q or R⁴ is hydroxy, mercapto or amino as appropriate;
- (d) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is (1-6C)alkanoylamino or substituted (2-6C)alkanoylamino, the acylation of a compound of the Formula I wherein a substituent on Q or R⁴ is amino;
 - (e) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is (1-6C)alkanesulphonylamino, the reaction of a compound of the Formula I wherein a substituent on Q or R⁴ is amino with a (1-6C)alkanesulphonic acid, or an activated derivative thereof;
- 20 (f) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is carboxy, carboxy-(1-6C)alkyl, carboxy-(1-6C)alkoxy, carboxy-(1-6C)alkylamino,

 N-(1-6C)alkyl-carboxy-(1-6C)alkylamino or carboxy-(2-6C)alkanoylamino, the cleavage of a compound of the Formula I wherein a substituent on Q or R⁴ is (1-6C)alkoxycarbonyl,

 (1-6C)alkoxycarbonyl-(1-6C)alkyl, (1-6C)alkoxycarbonyl-(1-6C)alkoxy,
- 25 (1-6C)alkoxycarbonyl-(1-6C)alkylamino, N-(1-6C)alkyl-(1-6C)alkòxycarbonyl-(1-6C)alkylamino or (1-6C)alkoxycarbonyl-(2-6C)alkanoylamino as appropriate;
 - (g) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is



amino-(1-6C)alkyl, heterocyclyl-(1-6C)alkyl, (1-6C)alkylamino-(1-6C)alkyl, di-[(1-6C)alkyl]amino-(1-6C)alkyl, substituted (2-6C)alkylamino-(1-6C)alkyl or substituted N-(1-6C)alkyl-(2-6C)alkylamino-(1-6C)alkyl, the reaction of a compound of the Formula I wherein a substituent on Q or R⁴ is a group of the formula -(1-6C)alkylene-Z wherein Z is a displaceable group with an appropriate amine or heterocyclyl compound;

- (h) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is amino, heterocyclyl, (1-6C)alkylamino, di-[(1-6C)alkyl]amino, substituted (1-6C)alkylamino, substituted N-(1-6C)alkyl-(1-6C)alkylamino, substituted (2-6C)alkylamino or substituted N-(1-6C)alkyl-(2-6C)alkylamino, the reaction of a compound of the Formula I wherein a substituent on Q or R⁴ is a displaceable group Z with an appropriate amine or heterocyclyl compound;
- (i) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is N-(1-6C)alkyl-(1-6C)alkanesulphonylamino, the alkylation, conveniently in the presence of a suitable base, of an amide derivative of the Formula I wherein a substituent on Q or R⁴ is
 15 (1-6C)alkanesulphonylamino;
- (j) for the preparation of a compound of the Formula I wherein a substituent on Q or R⁴ is a hydroxy-heterocyclyl-(1-6C)alkoxy group, a hydroxy-(1-6C)alkylamino-(2-6C)alkoxy group or a hydroxy-di-[(1-6C)alkyl]amino-(2-6C)alkoxy group, the reaction of a compound of the Formula I wherein a substituent on Q or R⁴ is a epoxy-substituted (1-6C)alkoxy group with a heterocyclyl compound or an appropriate arrine; or
 - (k) for the preparation of a compound of the Formula I wherein R^2 or a substituent on Q or R^4 is an amino group, the reduction of a compound of the Formula I wherein R^2 or a substituent on Q or R^4 is a nitro group.
- 25 10. A pharmaceutical composition which comprises an amide derivative of the Formula I, or a pharmaceutically-acceptable or <u>in-vivo</u>-cleavable ester thereof, according to claim 1 in association with a pharmaceutically-acceptable diluent or carrier.
- 11. The use of an amide derivative of the Formula I, or a pharmaceutically-acceptable salt or in-vivo-cleavable ester thereof, according to claim 1 in the manufacture of a medicament for use in the treatment of medical conditions mediated by cytokines.

